

=> fil reg

FILE 'REGISTRY' ENTERED AT 15:25:50 ON 06 MAY 2004  
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STRUCTURE FILE UPDATES: 5 MAY 2004 HIGHEST RN 680179-46-8  
 DICTIONARY FILE UPDATES: 5 MAY 2004 HIGHEST RN 680179-46-8

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 6, 2004

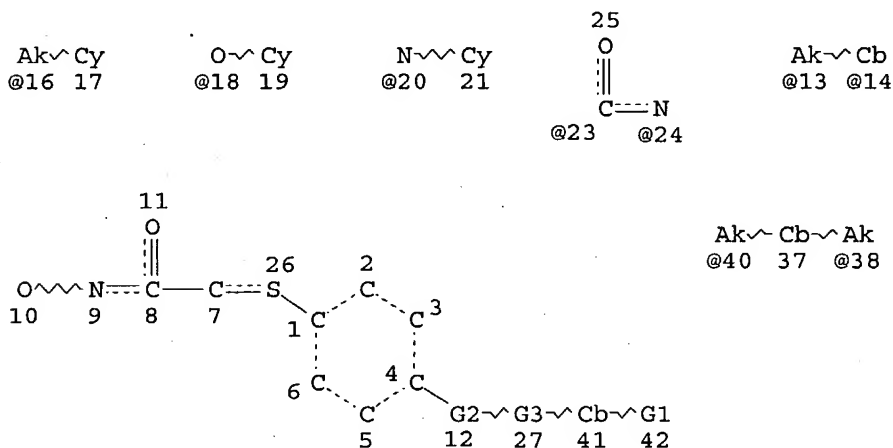
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Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more  
 information enter HELP PROP at an arrow prompt in the file or refer  
 to the file summary sheet on the web at:  
<http://www.cas.org/ONLINE/DBSS/registryss.html>

=> d sta que l7

L5 STR



VAR G1=CY/16/18/20  
 VAR G2=O/S/N/23-4 24-27/24-4 23-27/C  
 VAR G3=AK/CB/13-12 14-41/14-12 13-41/40-12 38-41  
 NODE ATTRIBUTES:  
 NSPEC IS R AT 7  
 DEFAULT MLEVEL IS ATOM  
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:  
 RING(S) ARE ISOLATED OR EMBEDDED  
 NUMBER OF NODES IS 30

STEREO ATTRIBUTES: NONE  
 L7 157 SEA FILE=REGISTRY SSS FUL L5

100.0% PROCESSED 3605 ITERATIONS  
 SEARCH TIME: 00.00.01

157 ANSWERS

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L3 STR L1  
L4 10 S L3  
L5 STR L3  
L6 10 S L5  
L7 157 S L5 FUL  
SAV L7 ZINNA657/A

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L8 0 S L7

FILE 'HCAPLUS' ENTERED AT 15:23:28 ON 06 MAY 2004

L9 2 S L7  
L10 1 S L9 AND PHARMACIA?/PA,CS  
L11 2 S L9 AND (FRESKOS ? OR FOBIAN ? OR BARTA ? OR BECKER ? OR BEDEL  
L12 2 S L9-L11

FILE 'USPATFULL, USPAT2' ENTERED AT 15:25:18 ON 06 MAY 2004

L13 3 S L7

FILE 'REGISTRY' ENTERED AT 15:25:50 ON 06 MAY 2004

=> fil uspatall

FILE 'USPATFULL' ENTERED AT 15:25:58 ON 06 MAY 2004  
CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'USPAT2' ENTERED AT 15:25:58 ON 06 MAY 2004  
CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

=>

=> d l13 bib abs hitrn fhitrstr tot

L13 ANSWER 1 OF 3 USPATFULL on STN

AN 2004:31882 USPATFULL

TI Aromatic sulfone hydroxamates and their use as protease inhibitors

IN Freskos, John N., Clayton, MO, UNITED STATES  
Fobian, Y vette M., Wildwood, MO, UNITED STATES  
Awasthi, Alok K., Skokie, IL, UNITED STATES  
Barta, Thomas E., Evanston, IL, UNITED STATES  
Becker, Daniel P., Glenview, IL, UNITED STATES  
Bedell, Louis J., Mt. Prospect, IL, UNITED STATES  
Boehm, Terri L., Ballwin, MO, UNITED STATES  
Carroll, Jeffery N., Columbia, IL, UNITED STATES  
Chandrakumar, Nizal S., Vernon Hills, IL, UNITED STATES  
DeCrescenzo, Gary A., St. Charles, MO, UNITED STATES  
Desai, Bipin N., Vernon Hills, IL, UNITED STATES  
Heron, Marcia I., Wester Springs, IL, UNITED STATES  
Hockerman, Susan L., Lincolnwood, IL, UNITED STATES  
Jull, Sara M., Villa Park, IL, UNITED STATES  
Kassab, Darren J., O' Fallon, MO, UNITED STATES  
Kolodziej, Steve A., Ballwin, MO, UNITED STATES  
McDonald, Joseph, Wildwood, MO, UNITED STATES  
Mischke, Deborah A., Defiance, MO, UNITED STATES  
Mullins, Patrick B., St Louis, MO, UNITED STATES  
Norton, Monica B., St. Louis, MO, UNITED STATES

Rico, Joseph G., Ballwin, MO, UNITED STATES  
Talley, John J., Cambridge, MA, UNITED STATES  
Trivedi, Mahima, Skokie, IL, UNITED STATES  
Villamil, Clara I., Glenview, IL, UNITED STATES  
Wang, Lijuan Jane, Wildwood, MO, UNITED STATES

PI US 2004024024 A1 20040205  
AI US 2002-291983 A1 20021112 (10)  
RLI Continuation-in-part of Ser. No. US 2002-142737, filed on 10 May 2002,  
PENDING  
PRAI US 2001-290375P 20010511 (60)  
DT Utility  
FS APPLICATION  
LREP HARNESS, DICKEY, & PIERCE, P.L.C, 7700 BONHOMME, STE 400, ST. LOUIS, MO,  
63105  
CLMN Number of Claims: 26  
ECL Exemplary Claim: 1  
DRWN No Drawings  
LN.CNT 11028

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB This invention is directed to aromatic sulfone hydroxamates (also known as "aromatic sulfone hydroxamic acids") and salts thereof that, inter alia, inhibit matrix metalloproteinase (also known as "matrix metalloprotease" or "MMP") activity and/or aggrecanase activity. This invention also is directed to a prevention or treatment method that comprises administering such a compound or salt in an MMP-inhibiting and/or aggrecanase-inhibiting effective amount to an animal, particularly a mammal having (or disposed to having) a pathological condition associated with MMP and/or aggrecanase activity.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 476182-38-4P 476182-39-5P 476182-40-8P  
476182-41-9P 476182-42-0P 476182-52-2P  
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476187-45-8P 476187-46-9P 476187-47-0P

476187-48-1P 476187-49-2P 476187-50-5P

476189-11-4P

(claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 476189-30-7P 476189-32-9P 476189-38-5P

476189-41-0P 476189-44-3P 476189-53-4P

476191-01-2P 476191-13-6P 476191-15-8P

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476195-24-1P 476195-25-2P 476195-26-3P

476195-27-4P 476195-31-0P 476195-33-2P

476195-41-2P

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 476189-73-8 476189-81-8 476189-91-0

476189-93-2 476189-95-4 476190-30-4

476191-35-2 476191-73-8 476191-87-4

476191-94-3 476192-10-6 476194-89-5

476195-23-0

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 476193-41-6P 476193-49-4P 476193-57-4P

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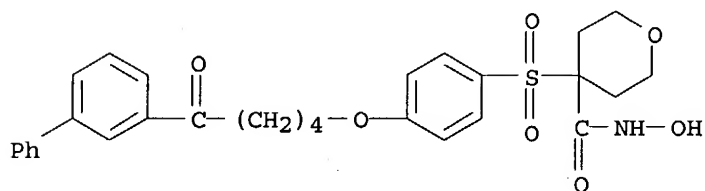
(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 476182-38-4P

(claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

RN 476182-38-4 USPATFULL

CN 2H-Pyran-4-carboxamide, 4-[[4-[(5-[1,1'-biphenyl]-3-yl-5-oxopentyl)oxy]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)



L13 ANSWER 2 OF 3 USPATFULL on STN

AN 2004:13496 USPATFULL

TI Aromatic sulfone hydroxamates and their use as protease inhibitors

IN Freskos, John N., Clayton, MO, UNITED STATES

Fobian, Yvette M., Wildwood, MO, UNITED STATES

Barta, Thomas E., Evanston, IL, UNITED STATES

Becker, Daniel P., Glenview, IL, UNITED STATES

Bedell, Louis J., Mt. Prospect, IL, UNITED STATES

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Hockerman, Susan L., Chicago, IL, UNITED STATES

Kassab, Darren J., Wildwood, MO, UNITED STATES

Kolodziej, Steve A., Ballwin, MO, UNITED STATES

McDonald, Joseph, Wildwood, MO, UNITED STATES

Mischke, Deborah A., Defiance, MO, UNITED STATES  
Norton, Monica B., St. Louis, MO, UNITED STATES  
Rico, Joseph G., Ballwin, MO, UNITED STATES  
Talley, John J., Cambridge, MA, UNITED STATES  
Villamil, Clara I., Glenview, IL, UNITED STATES  
Wang, Lijuan Jane, Wildwood, MO, UNITED STATES

PI US 2004010019 A1 20040115  
US 6689794 B2 20040210

AI US 2002-142737 A1 20020510 (10)

PRAI US 2001-290375P 20010511 (60)

DT Utility

FS APPLICATION

LREP David M. Gryte, Harness, Dickey & Pierce, P.L.C., Suite 400, 7700  
Bonhomme, St. Louis, MO, 63105

CLMN Number of Claims: 393

ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 15379

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB This invention is directed to aromatic sulfone hydroxamates (also known as "aromatic sulfone hydroxamic acids") and salts thereof that, inter alia, inhibit matrix metalloproteinase (also known as "matrix metalloprotease" or "MMP") activity and/or aggrecanase activity. This invention also is directed to a prevention or treatment method that comprises administering such a compound or salt in an MMP-inhibiting and/or aggrecanase-inhibiting effective amount to an animal, particularly a mammal having (or disposed to having) a pathological condition associated with MMP and/or aggrecanase activity.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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476182-41-9P 476182-42-0P 476182-52-2P  
476186-38-6P 476186-39-7P 476186-40-0P  
476186-41-1P 476186-42-2P 476186-43-3P  
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476187-45-8P 476187-46-9P 476187-47-0P  
 476187-48-1P 476187-49-2P 476187-50-5P  
 476189-11-4P

(claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 476189-30-7P 476189-32-9P 476189-38-5P  
 476189-41-0P 476189-44-3P 476189-53-4P  
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 476195-24-1P 476195-25-2P 476195-26-3P  
 476195-27-4P 476195-31-0P 476195-33-2P  
 476195-41-2P

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 476189-73-8 476189-81-8 476189-91-0  
 476189-93-2 476189-95-4 476190-30-4  
 476191-35-2 476191-73-8 476191-87-4  
 476191-94-3 476192-10-6 476194-89-5  
 476195-23-0

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

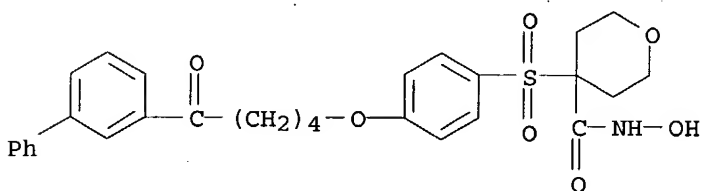
IT 476193-41-6P 476193-49-4P 476193-57-4P  
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 476194-29-3P 476194-38-4P

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 476182-38-4P  
 (claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

RN 476182-38-4 USPATFULL

CN 2H-Pyran-4-carboxamide, 4-[[4-[(5-[1,1'-biphenyl]-3-yl-5-oxopentyl)oxy]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)



L13 ANSWER 3 OF 3 USPAT2 on STN

AN 2004:13496 USPAT2

TI Aromatic sulfone hydroxamates and their use as protease inhibitors

IN Freskos, John N., Clayton, MO, United States  
 Fobian, Yvette M., Wildwood, MO, United States  
 Barta, Thomas E., Evanston, IL, United States  
 Becker, Daniel P., Glenview, IL, United States  
 Bedell, Louis J., Mt. Prospect, IL, United States  
 Boehm, Terri L., Ballwin, MO, United States  
 Carroll, Jeffery N., Columbia, IL, United States  
 DeCrescenzo, Gary A., St. Charles, MO, United States  
 Hockerman, Susan L., Chicago, IL, United States  
 Kassab, Darren J., Wildwood, MO, United States  
 Kolodziej, Steve A., Ballwin, MO, United States  
 McDonald, Joseph, Wildwood, MO, United States

Mischke, Deborah A., Defiance, MO, United States  
Norton, Monica B., St. Louis, MO, United States  
Rico, Joseph G., Ballwin, MO, United States  
Talley, John J., Boston, MA, United States  
Villamil, Clara I., Glenview, IL, United States  
Wang, Lijuan Jane, Wildwood, MO, United States  
PA Pharmacia Corporation, St. Louis, MO, United States (U.S. corporation)  
PI US 6689794 B2 20040210  
AI US 2002-142737 20020510 (10)  
PRAI US 2001-290375P 20010511 (60)  
DT Utility  
FS GRANTED  
EXNAM Primary Examiner: Davis, Zinna Northington  
LREP Harness, Dickey & Pierce, P.L.C.  
CLMN Number of Claims: 90  
ECL Exemplary Claim: 1  
DRWN 0 Drawing Figure(s); 0 Drawing Page(s)  
LN.CNT 9810  
CAS INDEXING IS AVAILABLE FOR THIS PATENT.  
AB This invention is directed to aromatic sulfone hydroxamates (also known as "aromatic sulfone hydroxamic acids") and salts thereof that, inter alia, inhibit matrix metalloproteinase (also known as "matrix metalloprotease" or "MMP") activity and/or aggrecanase activity. This invention also is directed to a prevention or treatment method that comprises administering such a compound or salt in an MMP-inhibiting and/or aggrecanase-inhibiting effective amount to an animal, particularly a mammal having (or disposed to having) a pathological condition associated with MMP and/or aggrecanase activity.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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476182-41-9P 476182-42-0P 476182-52-2P  
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 476189-11-4P

(claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 476189-30-7P 476189-32-9P 476189-38-5P  
 476189-41-0P 476189-44-3P 476189-53-4P  
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(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 476189-73-8 476189-81-8 476189-91-0  
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(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

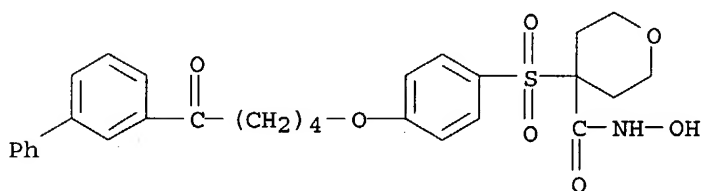
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 476194-29-3P 476194-38-4P

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 476182-38-4P  
 (claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

RN 476182-38-4 USPAT2

CN 2H-Pyran-4-carboxamide, 4-[[4-[(5-[1,1'-biphenyl]-3-yl-5-oxopentyl)oxy]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)



=> fil hcaplus

FILE 'HCAPLUS' ENTERED AT 15:26:41 ON 06 MAY 2004

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

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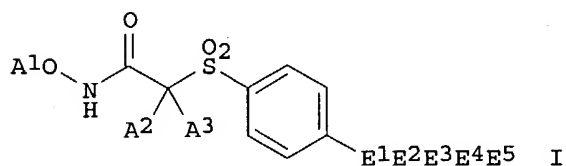
FILE COVERS 1907 - 6 May 2004 VOL 140 ISS 19  
FILE LAST UPDATED: 5 May 2004 (20040505/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

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L12 ANSWER 1 OF 2 HCAPLUS COPYRIGHT 2004 ACS on STN  
AN 2004:100823 HCAPLUS  
DN 140:163704  
ED Entered STN: 08 Feb 2004  
TI Preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors  
IN Freskos, John N.; Fobian, Yvette M.; Awasthi, Alok K.; Barta, Thomas E.; Becker, Daniel P.; Bedell, Louis J.; Boehm, Terri L.; Carroll, Jeffery N.; Chandrakumar, Nizal S.; Decrescenzo, Gary A.; Desai, Bipin N.; Heron, Marcia I.; Hockerman, Susan L.; Jull, Sara M.; Kassab, Darren J.; Kolodziej, Steve A.; McDonald, Joseph; Mischke, Deborah A.; Mullins, Patrick B.; Norton, Monica B.; Rico, Joseph G.; Talley, John J.; Trivedi, Mahima; Villamil, Clara I.; Wang, Lijuan Jane  
PA USA  
SO U.S. Pat. Appl. Publ., 365 pp., Cont.-in-part of U.S. Ser. No. 142,737. CODEN: USXXCO  
DT Patent  
LA English  
IC ICM C07D405-02  
ICS C07D403-02; A61K031-451; A61K031-454; A61K031-415  
NCL 514326000; 514513000; 514357000; 514408000; 514575000; 514382000; 514459000; 546210000; 546207000; 548252000  
CC 27-11 (Heterocyclic Compounds (One Hetero Atom))  
Section cross-reference(s): 1  
FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2004024024	A1	20040205	US 2002-291983	20021112
	US 2004010019	A1	20040115	US 2002-142737	20020510
	US 6689794	B2	20040210		
PRAI	US 2001-290375P	P	20010511		
	US 2002-142737	A2	20020510		
OS	MARPAT 140:163704				
GI					



AB Title compds. [I; A1 = H, (substituted) alkylcarbonyl, alkoxy carbonyl, carbocyclylcarbonyl, heterocyclylcarbonyl, aminoalkylthiocarbonyl, etc.; A2A3C = (substituted) heterocyclyl; E1 = O, S, SO, SO2, NR1, CONR1, CR1R2;

E2 = (substituted) alkyl, cycloalkyl, alkylcycloalkyl, cycloalkylalkyl, alkylcycloalkylalkyl; E3 = CO, O2C, CNR3, NR4, NR4SO2, S, SO, etc.; E4 = bond, (substituted) alkyl, alkenyl; E5 = H, OH, (substituted) alkyl, alkenyl, alkynyl, alkoxy, alkoxyalkyl, carbocyclyl, heterocyclyl; R1, R2 = H, (substituted) alkyl; R4 = H, alkyl, cycloalkyl, etc.; with provisos], were prepared Thus, tetrahydro-4-[[4-[[5-(4-methoxyphenyl)-5-oxopentyl]oxy]phenyl]sulfonyl]-2H-pyran-4-carboxylic acid 1,1-dimethylethyl ester (preparation given) in CH2Cl2 was treated with Me3SiCN and ZnI2 to give 81% cyanohydrin. The product in DMF was treated with 1-hydroxybenzotriazole, 1-(3-dimethylaminopropyl)-3-ethylcarbodiimide hydrochloride, N-methylmorpholine, and tetrahydropyranhydroxylamine to give 70% THP-protected hydroxamate. The latter was stirred with aqueous HCl in dioxane/MeOH to give 59% 4-[[4-[[[(4Z)-5-cyano-5-(4-methoxyphenyl)-4-pentenyl]oxy]phenyl]sulfonyl]tetrahydro-N-hydroxy-2H-pyran-4-carboxamide. This inhibited MMP-13 with IC50 = 0.2 nM.

- ST arylsulfonylpyranhydroxamate prepn matrix metalloprotease aggrecanase inhibitor; pyranhydroxamate arylsulfonyl prepn matrix metalloprotease aggrecanase inhibitor; arylsulfonylpyran hydroxamate prepn matrix metalloprotease aggrecanase inhibitor
- IT Nervous system, disease  
(central, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Nervous system, disease  
(degeneration, treatment of; preparation of arylsulfonylpyranhydroxamates for treating a pathol. condition of the CNS associated with nitrosative or oxidative stress)
- IT Brain, disease  
(ischemia, treatment of; preparation of arylsulfonylpyranhydroxamates for treating a pathol. condition of the CNS associated with nitrosative or oxidative stress)
- IT Human  
Nervous system agents  
(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Hydroxamic acids  
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Anti-ischemic agents  
Oxidative stress, biological  
(preparation of arylsulfonylpyranhydroxamates for treating a pathol. condition of the CNS associated with nitrosative or oxidative stress)
- IT Brain, disease  
(stroke, treatment of; preparation of arylsulfonylpyranhydroxamates for treating a pathol. condition of the CNS associated with nitrosative or oxidative stress)
- IT 308829-55-2P 476182-04-4P 476182-05-5P 476182-07-7P 476182-08-8P  
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RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU  
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES  
(Uses)

(claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix  
metalloprotease and/or aggrecanase inhibitors)

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	476184-94-8P	476184-95-9P	476184-96-0P	476184-97-1P	476184-98-2P
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476187-03-8P 476187-04-9P 476187-05-0P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU  
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES  
(Uses)

(claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix  
metalloprotease and/or aggrecanase inhibitors)

IT 476187-06-1P 476187-07-2P 476187-08-3P  
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RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU  
 (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES  
 (Uses)

(claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix  
 metalloprotease and/or aggrecanase inhibitors)

IT 146480-35-5, Matrix metalloproteinase-2 146480-36-6, Matrix  
 metalloproteinase-9 147172-61-0, Aggrecanase 175449-82-8, Matrix  
 metalloproteinase-13

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (inhibitors; preparation of arylsulfonylpyranhydroxamates as matrix  
 metalloprotease and/or aggrecanase inhibitors)

IT 151769-16-3, Tnf- $\alpha$  convertase

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (pathol. condition treatment; preparation of arylsulfonylpyranhydroxamates  
 as matrix metalloprotease and/or aggrecanase inhibitors)

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 654639-26-6P 654639-27-7P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU

(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 476189-58-9 476189-67-0 476189-70-5 476189-73-8  
 476189-76-1 476189-79-4 476189-81-8 476189-87-4  
 476189-89-6 476189-91-0 476189-93-2  
 476189-95-4 476189-97-6 476189-99-8 476190-01-9  
 476190-03-1 476190-05-3 476190-07-5 476190-11-1 476190-14-4  
 476190-16-6 476190-18-8 476190-20-2 476190-22-4 476190-26-8  
 476190-28-0 476190-30-4 476190-32-6 476190-34-8  
 476190-36-0 476190-38-2 476190-39-3 476190-41-7 476190-73-5  
 476190-75-7 476190-81-5 476190-83-7 476190-89-3 476190-91-7  
 476190-93-9 476191-03-4 476191-05-6 476191-07-8 476191-09-0  
 476191-35-2 476191-37-4 476191-39-6 476191-41-0  
 476191-43-2 476191-45-4 476191-47-6 476191-49-8 476191-51-2  
 476191-55-6 476191-73-8 476191-75-0 476191-81-8  
 476191-87-4 476191-94-3 476191-96-5  
 476192-10-6 476194-86-2 476194-87-3 476194-89-5  
 476194-96-4 476194-97-5 476194-98-6 476194-99-7 476195-00-3  
 476195-01-4 476195-02-5 476195-03-6 476195-04-7 476195-05-8  
 476195-06-9 476195-14-9 476195-17-2 476195-23-0  
 476195-38-7 476195-39-8

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 75-03-6, Iodoethane 75-16-1, Methylmagnesium bromide 91-21-4,  
 1,2,3,4-Tetrahydroisoquinoline 95-55-6, 2-Aminophenol 96-34-4, Methyl  
 chloroacetate 96-48-0,  $\gamma$ -Butyrolactone 98-80-6, Phenylboronic  
 acid 100-07-2, p-Anisoyl chloride 104-92-7, 4-Bromoanisole 106-53-6,  
 4-Bromobenzenethiol 107-18-6, Allyl alcohol, reactions 107-59-5,  
 tert-Butyl chloroacetate 124-63-0, Methylsulfonyl chloride 332-25-2,  
 4-Trifluoromethoxybenzonitrile 371-42-6, 4-Fluorothiophenol 580-13-2,  
 2-Bromonaphthalene 586-77-6, 4-Bromo-N,N-dimethylaniline 821-09-0,  
 4-Penten-1-ol 867-13-0, Triethyl phosphonoacetate 873-62-1,  
 3-Cyanophenol 883-44-3, N-(3-Hydroxypropyl)phthalimide 1126-09-6,  
 Ethyl isonipecotate 1679-18-1, 4-Chlorobenzeneboronic acid 1765-93-1,  
 4-Fluorobenzeneboronic acid 2382-96-9, 2-Mercaptobenzoxazole  
 3161-51-1, 3-(Dibenzylamino)-1-propanol 4799-68-2, 3-Benzoyloxy-1-  
 propanol 5292-43-3, tert-Butyl bromoacetate 5414-19-7,  
 Bis(2-bromoethyl)ether 6482-24-2, 2-Bromoethyl methyl ether 7051-34-5,  
 Bromomethylcyclopropane 7658-80-2, o-Toluic hydrazide 17715-69-4,  
 1-Bromo-2,4-dimethoxybenzene 17997-47-6, 2-Tributylstannylpyridine  
 18791-75-8, 4-Bromo-2-thiophenecarboxaldehyde 27374-25-0,  
 [(1-Ethoxycyclopropyl)oxy]trimethylsilane 28229-69-8, 3-Bromophenethyl  
 alcohol 42287-90-1 42330-88-1, 2-(3-Chloropropoxy)tetrahydro-2H-pyran  
 52898-32-5, N-(3-Buten-1-yl)phthalimide 55162-82-8, 5-Benzoyloxy-1-  
 pentanol 56935-71-8, 4-(Trifluoromethoxy)benzamidoxime 58885-58-8,  
 tert-Butyl N-(3-hydroxypropyl)carbamate 89691-67-8, 2-Bromo-4-  
 methoxyacetophenone 144025-03-6, 2,4-Difluorophenylboronic acid  
 155288-39-4 168267-41-2, 3,4-Difluorobenzeneboronic acid 226396-34-5,  
 Ethyl 4-[(4-fluorophenyl)sulfonyl]-1-(2-methoxyethyl)piperidine-4-  
 carboxylate 476194-58-8 476194-61-3

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT 99-98-9P, N,N-Dimethyl-1,4-phenylenediamine 405-31-2P 65537-54-4P  
 71912-71-5P, [1,1'-Biphenyl]-3-ethanol 136416-19-8P,  
 [1,1'-Biphenyl]-3-propanol 142851-03-4P 212770-40-6P 212770-41-7P  
 226389-21-5P 226396-33-4P, Ethyl 4-[(4-fluorophenyl)sulfonyl]piperidine-  
 4-carboxylate hydrochloride 226396-62-9P 226396-63-0P 226396-70-9P  
 226396-71-0P 226396-72-1P 226398-02-3P 226399-90-2P 226401-27-0P

283153-83-3P 476189-15-8P 476189-17-0P 476189-19-2P 476189-20-5P  
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 476192-17-3P 476192-19-5P 476192-21-9P 476192-23-1P 476192-25-3P  
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 476193-39-2P **476193-41-6P** 476193-43-8P 476193-45-0P  
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 476193-55-2P **476193-57-4P** 476193-60-9P 476193-62-1P  
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**476193-90-5P** 476193-92-7P 476193-94-9P **476193-95-0P**  
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 476194-67-9P 476194-68-0P 476194-70-4P 476194-71-5P 476194-72-6P  
 476194-73-7P 476194-75-9P 476195-43-4P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

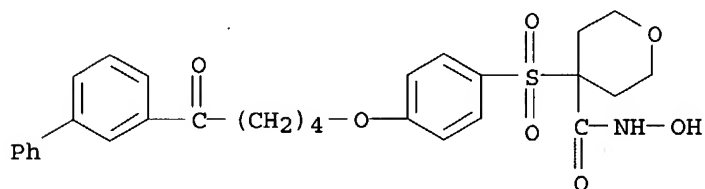
IT **476182-38-4P**

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

RN 476182-38-4 HCAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(5-[1,1'-biphenyl]-3-yl-5-oxopentyl)oxy]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)



L12 ANSWER 2 OF 2 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2002:888730 HCAPLUS

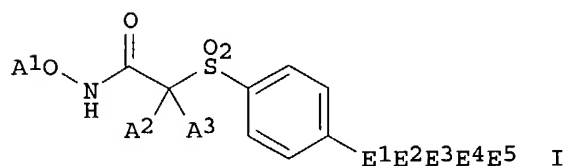
DN 137:384747

ED Entered STN: 22 Nov 2002

TI Preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors

IN Barta, Thomas E.; Becker, Daniel P.; Bedell,  
Louis J.; Boehm, Terri L.; Fobian, Yvette M.;  
Freskos, John N.; Hockerman, Susan L.; Kassab,  
Darren J.; Kolodziej, Steve A.; McDonald, Joseph J.  
; Norton, Monica B.; Rico, Joseph G.; Talley,  
John J.; Villamil, Clara I.; Wang, Tijuana Jane  
PA Pharmacia Corporation, USA  
SO PCT Int. Appl., 627 pp.  
CODEN: PIXXD2  
DT Patent  
LA English  
IC ICM C07D309-08  
ICS C07D405-12; C07D401-12; C07D211-66; C07D407-12; C07D417-12;  
C07D409-12; C07D413-12; A61K031-351; A61P035-00  
CC 27-11 (Heterocyclic Compounds (One Hetero Atom))  
Section cross-reference(s): 1  
FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002092588	A2	20021121	WO 2002-US15257	20020510
	WO 2002092588	A3	20030227		
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
EP	1385836	A2	20040204	EP 2002-729204	20020510
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR			
BR	2002009525	A	20040309	BR 2002-9525	20020510
NO	2003004995	A	20031216	NO 2003-4995	20031110
PRAI	US 2001-290375P	P	20010511		
	WO 2002-US15257	W	20020510		
OS	MARPAT 137:384747				
GI					



AB Title compds. [I; A1 = H, (substituted) alkylcarbonyl, alkoxy carbonyl, carbocyclylcarbonyl, heterocyclylcarbonyl, aminoalkylthiocarbonyl, etc.; A2A3C = (substituted) heterocyclyl; E1 = O, S, SO, SO2, NR1, CONR1, CR1R2; E2 = (substituted) alkyl, cycloalkyl, alkylcycloalkyl, cycloalkylalkyl, alkylcycloalkylalkyl; E3 = CO, O2C, CNR3, NR4, NR4SO2, S, SO, etc.; E4 = bond, (substituted) alkyl, alkenyl; E5 = H, OH, (substituted) alkyl, alkenyl, alkynyl, alkoxy, alkoxyalkyl, carbocyclyl, heterocyclyl; R1, R2 = H, (substituted) alkyl; with provisos], were prepared Thus, tetrahydro-4-[[4-[[5-(4-methoxyphenyl)-5-oxopentyl]oxy]phenyl]sulfonyl]-2H-pyran-4-carboxylic acid 1,1-dimethylethyl ester (preparation given) in CH2Cl2



was treated with Me<sub>3</sub>SiCN and ZnI<sub>2</sub> to give 81% cyanohydrin. The product in DMF was treated with 1-hydroxybenzotriazole, 1-(3-dimethylaminopropyl)-3-ethylcarbodiimide hydrochloride, N-methylmorpholine, and tetrahydropyranhydroxylamine to give 70% THP-protected hydroxamate. The latter was stirred with aqueous HCl in dioxane/MeOH to give 59% 4-[[4-[[[(4Z)-5-cyano-5-(4-methoxyphenyl)-4-pentenyl]oxy]phenyl]sulfonyl]tetrahydro-N-hydroxy-2H-pyran-4-carboxamide. This inhibited MMP-13 with IC<sub>50</sub> = 0.2 nM.

- ST arylsulfonylpyranhydroxamate prepn matrix metalloprotease aggrecanase inhibitor; pyranhydroxamate arylsulfonyl prepn matrix metalloprotease aggrecanase inhibitor; drug arylsulfonylpyran hydroxamate prepn
- IT Antiarteriosclerotics  
(antiatherosclerotics; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Aneurysm  
(aortic, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Heart, disease  
(cardiomyopathy, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Nervous system, disease  
(central, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Lung, disease  
(chronic obstructive, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Eye, disease  
(cornea, ulcer, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Thrombosis  
(coronary arterial, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Artery, disease  
(coronary, thrombosis, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Radiation  
(damage, biol., treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Animal tissue  
(destruction treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Skin, disease  
(epidermolysis bullosa, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Heart, disease  
(failure, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Neoplasm  
(metastasis, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Infection  
(postmyocardial, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)
- IT Anti-Alzheimer's agents
- Anti-inflammatory agents
- Antiarthritics
- Anticoagulants
- Antipyretics
- Antitumor agents
- Human

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT Hydroxamic acids  
 RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT Proteins  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (proteinuria, treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT Alzheimer's disease  
 Anorexia  
 Arthritis  
 Atherosclerosis  
 Autoimmune disease  
 Bone, disease  
 Cachexia  
 Cardiovascular system, disease  
 Cirrhosis  
 Coagulation  
 Emphysema  
 Eye, disease  
 Fever and Hyperthermia  
 Fibrosis  
 Hemorrhage  
 Inflammation  
 Kidney, disease  
 Liver, disease  
 Lung, disease  
 Multiple sclerosis  
 Neoplasm  
 Osteoarthritis  
 Psoriasis  
 Rheumatoid arthritis  
 Sepsis  
 Shock (circulatory collapse)  
 Transplant rejection  
 Ulcer  
 (treatment; preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease and/or aggrecanase inhibitors)

IT

308829-55-2P	476182-04-4P	476182-05-5P	476182-07-7P	476182-08-8P
476182-09-9P	476182-10-2P	476182-11-3P	476182-12-4P	476182-13-5P
476182-14-6P	476182-15-7P	476182-16-8P	476182-17-9P	476182-18-0P
476182-19-1P	476182-20-4P	476182-21-5P	476182-22-6P	476182-23-7P
476182-24-8P	476182-25-9P	476182-26-0P	476182-27-1P	476182-28-2P
476182-29-3P	476182-30-6P	476182-32-8P	476182-35-1P	476182-36-2P
476182-37-3P	476182-38-4P	476182-39-5P		
476182-40-8P	476182-41-9P	476182-42-0P		
476182-43-1P	476182-44-2P	476182-45-3P	476182-46-4P	476182-47-5P
476182-48-6P	476182-50-0P	476182-52-2P	476182-53-3P	
476182-55-5P	476182-57-7P	476182-60-2P	476182-62-4P	476182-64-6P
476182-66-8P	476182-68-0P	476182-69-1P	476182-70-4P	476182-71-5P
476182-72-6P	476182-73-7P	476182-74-8P	476182-75-9P	476182-76-0P
476182-77-1P	476182-78-2P	476182-79-3P	476182-80-6P	476182-81-7P
476182-82-8P	476182-83-9P	476182-84-0P	476182-85-1P	476182-86-2P
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476182-93-1P	476182-94-2P	476182-95-3P	476182-97-5P	476182-98-6P
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476183-20-7P	476183-21-8P	476183-22-9P	476183-23-0P	476183-24-1P

476183-25-2P	476183-26-3P	476183-27-4P	476183-28-5P	476183-30-9P
476183-32-1P	476183-33-2P	476183-34-3P	476183-35-4P	476183-36-5P
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476183-42-3P	476183-44-5P	476183-45-6P	476183-46-7P	476183-47-8P
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476183-54-7P	476183-55-8P	476183-56-9P	476183-57-0P	476183-58-1P
476183-59-2P	476183-60-5P	476183-61-6P	476183-62-7P	476183-63-8P
476183-64-9P	476183-65-0P	476183-66-1P	476183-67-2P	476183-68-3P
476183-69-4P	476183-70-7P	476183-71-8P	476183-72-9P	476183-73-0P
476183-74-1P	476183-75-2P	476183-76-3P	476183-77-4P	476183-78-5P
476183-79-6P	476183-80-9P	476183-81-0P	476183-82-1P	476183-83-2P
476183-84-3P	476183-85-4P	476183-86-5P	476183-87-6P	476183-88-7P
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476184-14-2P	476184-15-3P	476184-16-4P	476184-17-5P	476184-18-6P
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476184-34-6P	476184-35-7P	476184-36-8P	476184-37-9P	476184-38-0P
476184-39-1P	476184-40-4P	476184-41-5P	476184-42-6P	476184-43-7P
476184-44-8P	476184-45-9P	476184-46-0P	476184-47-1P	476184-48-2P
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476184-59-5P	476184-60-8P	476184-61-9P	476184-62-0P	476184-63-1P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU  
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES  
(Uses)

(claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix  
metalloprotease and/or aggrecanase inhibitors)

IT	476184-64-2P	476184-65-3P	476184-66-4P	476184-67-5P	476184-68-6P
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	476184-74-4P	476184-75-5P	476184-76-6P	476184-77-7P	476184-78-8P
	476184-79-9P	476184-80-2P	476184-81-3P	476184-82-4P	476184-83-5P
	476184-84-6P	476184-85-7P	476184-86-8P	476184-87-9P	476184-88-0P
	476184-89-1P	476184-90-4P	476184-91-5P	476184-92-6P	476184-93-7P
	476184-94-8P	476184-95-9P	476184-96-0P	476184-97-1P	476184-98-2P
	476184-99-3P	476185-00-9P	476185-01-0P	476185-02-1P	476185-03-2P
	476185-04-3P	476185-05-4P	476185-06-5P	476185-07-6P	476185-08-7P
	476185-09-8P	476185-10-1P	476185-11-2P	476185-12-3P	476185-13-4P
	476185-14-5P	476185-15-6P	476185-16-7P	476185-17-8P	476185-18-9P
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	476185-37-2P	476185-38-3P	476185-40-7P	476185-42-9P	476185-44-1P
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	476185-70-3P	476185-71-4P	476185-72-5P	476185-73-6P	476185-74-7P
	476185-75-8P	476185-76-9P	476185-77-0P	476185-78-1P	476185-79-2P
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	476185-85-0P	476185-86-1P	476185-87-2P	476185-88-3P	476185-89-4P
	476185-90-7P	476185-91-8P	476185-92-9P	476185-93-0P	476185-94-1P
	476185-95-2P	476185-96-3P	476185-97-4P	476185-98-5P	476185-99-6P
	476186-00-2P	476186-01-3P	476186-02-4P	476186-03-5P	476186-04-6P
	476186-05-7P	476186-06-8P	476186-07-9P	476186-08-0P	476186-09-1P
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	476186-15-9P	476186-16-0P	476186-17-1P	476186-18-2P	476186-19-3P
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476187-03-8P 476187-04-9P 476187-05-0P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU  
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES  
(Uses)

(claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix  
metalloprotease and/or aggrecanase inhibitors)

IT 476187-06-1P 476187-07-2P 476187-08-3P  
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476187-94-7P 476187-95-8P 476187-96-9P 476187-97-0P 476187-98-1P  
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 476189-08-9P 476189-09-0P **476189-11-4P** 476189-13-6P  
 476195-35-4P 476195-36-5P 476195-37-6P 476195-40-1P 476195-42-3P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU  
 (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES  
 (Uses)

(claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix  
 metalloprotease and/or aggrecanase inhibitors)

IT 146480-35-5, Matrix metalloproteinase-2 146480-36-6, Matrix  
 metalloproteinase-9 147172-61-0, Aggrecanase 175449-82-8, Matrix  
 metalloproteinase-13

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (inhibitors; preparation of arylsulfonylpyranhydroxamates as matrix  
 metalloprotease and/or aggrecanase inhibitors)

IT 151769-16-3, Tnf- $\alpha$  convertase

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (pathol. condition treatment; preparation of arylsulfonylpyranhydroxamates  
 as matrix metalloprotease and/or aggrecanase inhibitors)

IT 476189-26-1P 476189-28-3P **476189-30-7P** **476189-32-9P**  
 476189-34-1P 476189-36-3P **476189-38-5P** **476189-41-0P**  
**476189-44-3P** 476189-47-6P 476189-49-8P 476189-51-2P  
**476189-53-4P** 476189-55-6P 476189-61-4P 476189-64-7P  
 476189-83-0P 476190-09-7P 476190-24-6P 476190-43-9P 476190-45-1P  
 476190-47-3P 476190-49-5P 476190-51-9P 476190-53-1P 476190-55-3P  
 476190-57-5P 476190-59-7P 476190-61-1P 476190-63-3P 476190-65-5P  
 476190-67-7P 476190-69-9P 476190-71-3P 476190-77-9P 476190-79-1P  
 476190-85-9P 476190-87-1P 476190-95-1P 476190-97-3P 476190-99-5P  
**476191-01-2P** 476191-11-4P **476191-13-6P**  
**476191-15-8P** **476191-17-0P** 476191-19-2P 476191-21-6P  
**476191-23-8P** **476191-25-0P** **476191-27-2P**  
**476191-29-4P** **476191-31-8P** **476191-33-0P**  
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 476191-98-7P 476192-00-4P 476192-02-6P **476192-04-8P**  
 476192-06-0P 476192-08-2P 476194-76-0P 476194-77-1P 476194-80-6P  
**476194-81-7P** 476194-82-8P **476194-83-9P** 476194-85-1P  
 476194-88-4P **476194-90-8P** 476194-91-9P 476194-92-0P  
 476194-93-1P 476194-94-2P 476195-11-6P 476195-12-7P 476195-15-0P  
 476195-16-1P 476195-18-3P 476195-20-7P 476195-21-8P  
**476195-22-9P** **476195-24-1P** **476195-25-2P**  
**476195-26-3P** **476195-27-4P** 476195-29-6P 476195-30-9P  
**476195-31-0P** **476195-33-2P** 476195-34-3P  
**476195-41-2P**

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU  
 (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES  
 (Uses)

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease  
 and/or aggrecanase inhibitors)

IT 476189-58-9 476189-67-0 476189-70-5 **476189-73-8**  
 476189-76-1 476189-79-4 **476189-81-8** 476189-87-4  
 476189-89-6 **476189-91-0** **476189-93-2**  
**476189-95-4** 476189-97-6 476189-99-8 476190-01-9  
 476190-03-1 476190-05-3 476190-07-5 476190-11-1 476190-14-4  
 476190-16-6 476190-18-8 476190-20-2 476190-22-4 476190-26-8  
 476190-28-0 **476190-30-4** 476190-32-6 476190-34-8  
 476190-36-0 476190-38-2 476190-39-3 476190-41-7 476190-73-5

476190-75-7 476190-81-5 476190-83-7 476190-89-3 476190-91-7  
 476190-93-9 476191-03-4 476191-05-6 476191-07-8 476191-09-0  
 476191-35-2 476191-37-4 476191-39-6 476191-41-0  
 476191-43-2 476191-45-4 476191-47-6 476191-49-8 476191-51-2  
 476191-55-6 476191-73-8 476191-75-0 476191-81-8  
 476191-87-4 476191-94-3 476191-96-5  
 476192-10-6 476194-86-2 476194-87-3 476194-89-5  
 476194-96-4 476194-97-5 476194-98-6 476194-99-7 476195-00-3  
 476195-01-4 476195-02-5 476195-03-6 476195-04-7 476195-05-8  
 476195-06-9 476195-14-9 476195-17-2 476195-23-0  
 476195-38-7 476195-39-8

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
 (Biological study); USES (Uses)

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease  
 and/or aggrecanase inhibitors)

IT 75-03-6, Iodoethane 75-16-1, Methylmagnesium bromide 91-21-4,  
 1,2,3,4-Tetrahydroisoquinoline 95-55-6, 2-Aminophenol 96-34-4, Methyl  
 chloroacetate 96-48-0,  $\gamma$ -Butyrolactone 98-80-6, Phenylboronic  
 acid 100-07-2, p-Anisoyl chloride 104-92-7, 4-Bromoanisole 106-53-6,  
 4-Bromobenzenethiol 107-18-6, Allyl alcohol, reactions 107-59-5,  
 tert-Butyl chloroacetate 124-63-0, Methylsulfonyl chloride 332-25-2,  
 4-Trifluoromethoxybenzonitrile 371-42-6, 4-Fluorothiophenol 580-13-2,  
 2-Bromonaphthalene 586-77-6, 4-Bromo-N,N-dimethylaniline 821-09-0,  
 4-Penten-1-ol 867-13-0, Triethyl phosphonoacetate 873-62-1,  
 3-Cyanophenol 883-44-3, N-(3-Hydroxypropyl)phthalimide 1126-09-6,  
 Ethyl isonipicotate 1679-18-1, 4-Chlorobenzeneboronic acid 1765-93-1,  
 4-Fluorobenzeneboronic acid 2382-96-9, 2-Mercaptobenzoxazole  
 3161-51-1, 3-(Dibenzylamino)-1-propanol 4799-68-2, 3-Benzoyloxy-1-  
 propanol 5292-43-3, tert-Butyl bromoacetate 5414-19-7,  
 Bis(2-bromoethyl)ether 6482-24-2, 2-Bromoethyl methyl ether 7051-34-5,  
 Bromomethylcyclopropane 7658-80-2, o-Toluic hydrazide 17715-69-4,  
 1-Bromo-2,4-dimethoxybenzene 17997-47-6, 2-Tributylstannylpyridine  
 18791-75-8, 4-Bromo-2-thiophenecarboxaldehyde 27374-25-0,  
 [(1-Ethoxycyclopropyl)oxy]trimethylsilane 28229-69-8, 3-Bromophenethyl  
 alcohol 42287-90-1 42330-88-1, 2-(3-Chloropropoxy)tetrahydro-2H-pyran  
 52898-32-5, N-(3-Buten-1-yl)phthalimide 55162-82-8, 5-Benzoyloxy-1-  
 pentanol 56935-71-8, 4-(Trifluoromethoxy)benzamidoxime 58885-58-8,  
 tert-Butyl N-(3-hydroxypropyl)carbamate 89691-67-8, 2-Bromo-4-  
 methoxyacetophenone 144025-03-6, 2,4-Difluorophenylboronic acid  
 155288-39-4 168267-41-2, 3,4-Difluorobenzeneboronic acid 226396-34-5,  
 Ethyl 4-[(4-fluorophenyl)sulfonyl]-1-(2-methoxyethyl)piperidine-4-  
 carboxylate 476194-58-8 476194-61-3

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease  
 and/or aggrecanase inhibitors)

IT 99-98-9P, N,N-Dimethyl-1,4-phenylenediamine 405-31-2P 65537-54-4P  
 71912-71-5P, [1,1'-Biphenyl]-3-ethanol 136416-19-8P,  
 [1,1'-Biphenyl]-3-propanol 142851-03-4P 212770-40-6P 212770-41-7P  
 226389-21-5P 226396-33-4P, Ethyl 4-[(4-fluorophenyl)sulfonyl]piperidine-  
 4-carboxylate hydrochloride 226396-62-9P 226396-63-0P 226396-70-9P  
 226396-71-0P 226396-72-1P 226398-02-3P 226399-90-2P 226401-27-0P  
 283153-83-3P 476189-15-8P 476189-17-0P 476189-19-2P 476189-20-5P  
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476193-27-8P 476193-29-0P 476193-31-4P 476193-33-6P 476193-36-9P  
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 476194-67-9P 476194-68-0P 476194-70-4P 476194-71-5P 476194-72-6P  
 476194-73-7P 476194-75-9P 476195-43-4P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT  
 (Reactant or reagent)  
 (preparation of arylsulfonylpyranhydroxamates as matrix metalloprotease  
 and/or aggrecanase inhibitors)

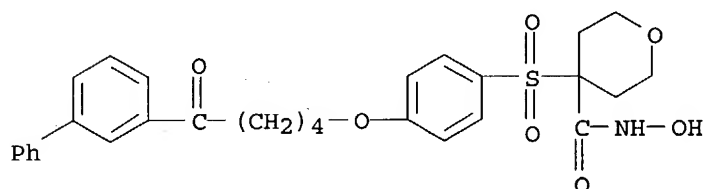
IT **476182-38-4P**

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU  
 (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES  
 (Uses)

(claimed compound; preparation of arylsulfonylpyranhydroxamates as matrix  
 metalloprotease and/or aggrecanase inhibitors)

RN 476182-38-4 HCAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(5-[1,1'-biphenyl]-3-yl-5-oxopentyl)oxy]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)



=> fil reg

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DICTIONARY FILE UPDATES: 5 MAY 2004 HIGHEST RN 680179-46-8

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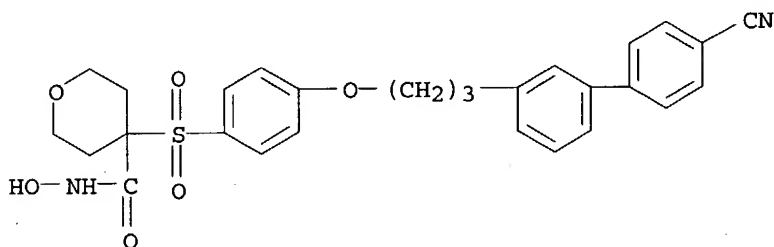
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 conducting SmartSELECT searches.

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<http://www.cas.org/ONLINE/DBSS/registryss.html>

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RN 476195-41-2 REGISTRY  
CN 2H-Pyran-4-carboxamide, 4-[[4-[3-(4'-cyano[1,1'-biphenyl]-3-yl)propoxy]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)  
FS 3D CONCORD  
MF C28 H28 N2 O6 S  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

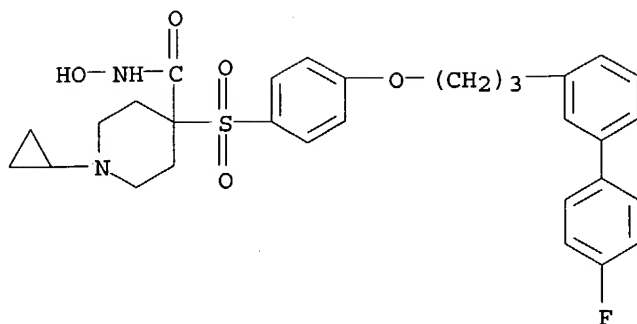
2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 10 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 476194-90-8 REGISTRY  
CN 4-Piperidinecarboxamide, 1-cyclopropyl-4-[[4-[3-(4'-fluoro[1,1'-biphenyl]-3-yl)propoxy]phenyl]sulfonyl]-N-hydroxy-, monohydrochloride (9CI) (CA INDEX NAME)  
MF C30 H33 F N2 O5 S . Cl H  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL  
CRN (476186-80-8)





● HCl

2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 20 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN

RN 476193-90-5 REGISTRY

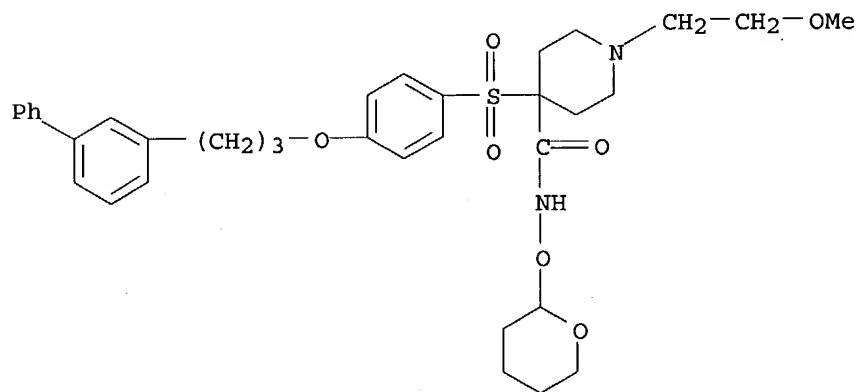
CN 4-Piperidinecarboxamide, 4-[[4-(3-[1,1'-biphenyl]-3-ylpropoxy)phenyl]sulfonyl]-1-(2-methoxyethyl)-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C35 H44 N2 O7 S

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



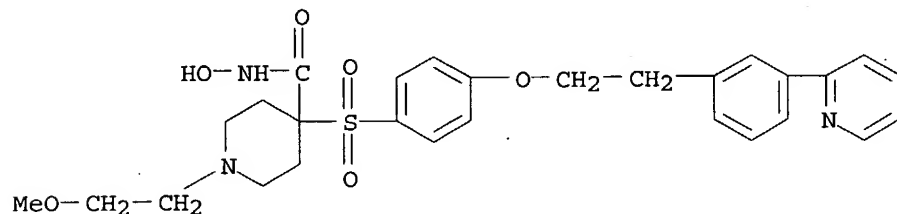
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2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 30 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
 RN 476191-33-0 REGISTRY  
 CN 4-Piperidinecarboxamide, N-hydroxy-1-(2-methoxyethyl)-4-[[4-[2-[3-(2-pyridinyl)phenyl]ethoxy]phenyl]sulfonyl]-, monohydrochloride (9CI) (CA INDEX NAME)  
 MF C28 H33 N3 O6 S . Cl H.  
 SR CA  
 LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL  
 CRN (476186-57-9)



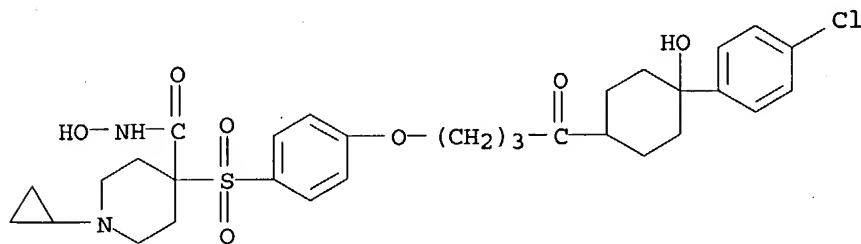
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2 REFERENCES IN FILE CA (1907 TO DATE)  
 2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 40 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
 RN 476190-30-4 REGISTRY  
 CN 4-Piperidinecarboxamide, 4-[[4-[4-[4-(4-chlorophenyl)-4-hydroxycyclohexyl]-4-oxobutoxy]phenyl]sulfonyl]-1-cyclopropyl-N-hydroxy-, monohydrochloride (9CI) (CA INDEX NAME)  
 MF C31 H39 Cl N2 O7 S . Cl H  
 SR CA  
 LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL  
 CRN (476182-52-2)



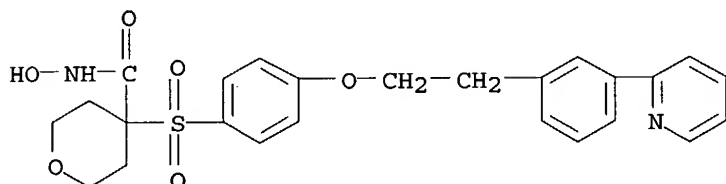
● HCl

2 REFERENCES IN FILE CA (1907 TO DATE)  
 2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 50 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 476189-32-9 REGISTRY  
CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[2-[3-(2-pyridinyl)phenyl]ethoxy]phenyl]sulfonyl]-, monohydrochloride (9CI) (CA INDEX NAME)  
MF C25 H26 N2 O6 S . Cl H  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL  
CRN (476186-55-7)



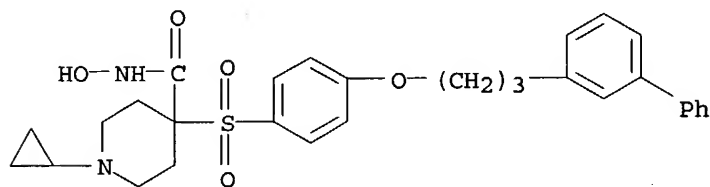
● HCl

2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 60 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 476187-42-5 REGISTRY  
CN 4-Piperidinecarboxamide, 4-[[4-(3-[1,1'-biphenyl]-3-ylpropoxy)phenyl]sulfonyl]-1-cyclopropyl-N-hydroxy- (9CI) (CA INDEX NAME)  
FS 3D CONCORD  
MF C30 H34 N2 O5 S  
CI COM  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



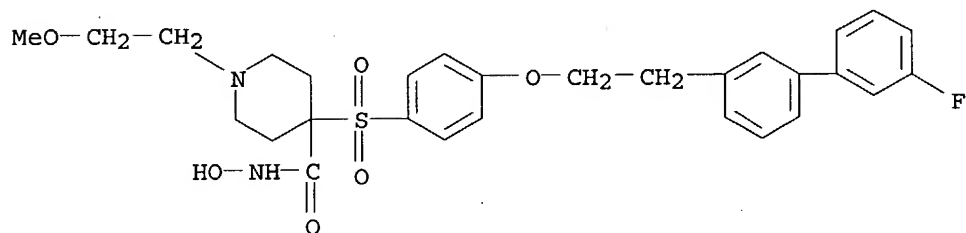
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2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 70 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 476187-28-7 REGISTRY  
CN 4-Piperidinecarboxamide, 4-[[4-[2-(3'-fluoro[1,1'-biphenyl]-3-yl)ethoxy]phenyl]sulfonyl]-N-hydroxy-1-(2-methoxyethyl)- (9CI) (CA INDEX NAME)  
FS 3D CONCORD  
MF C29 H33 F N2 O6 S  
CI COM  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



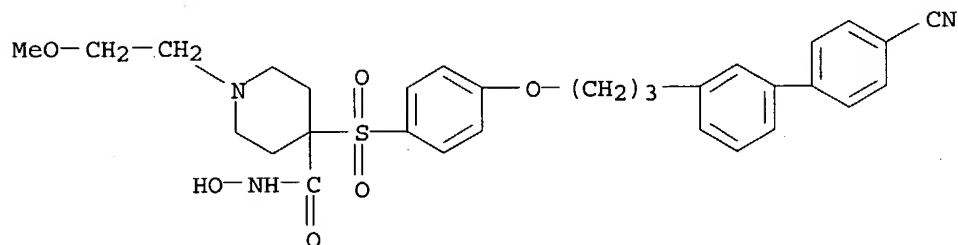
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2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 80 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 476187-11-8 REGISTRY  
CN 4-Piperidinecarboxamide, 4-[[4-[3-(4'-cyano[1,1'-biphenyl]-3-yl)propoxy]phenyl]sulfonyl]-N-hydroxy-1-(2-methoxyethyl)- (9CI) (CA INDEX NAME)  
FS 3D CONCORD  
MF C31 H35 N3 O6 S  
CI COM  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



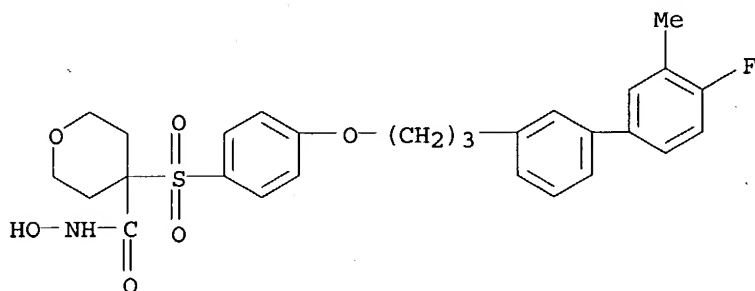
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2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 90 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 476187-00-5 REGISTRY  
CN 2H-Pyran-4-carboxamide, 4-[[4-[3-(4'-fluoro-3'-methyl[1,1'-biphenyl]-3-yl)propoxy]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)  
FS 3D CONCORD  
MF C28 H30 F N O6 S  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



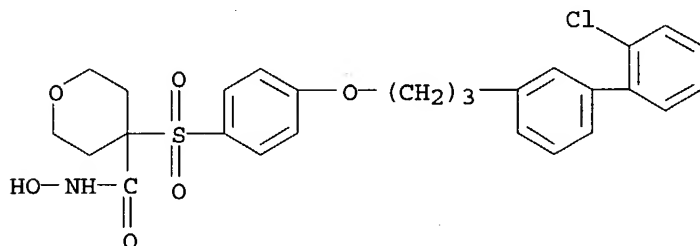
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2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 100 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 476186-89-7 REGISTRY  
CN 2H-Pyran-4-carboxamide, 4-[[4-[3-(2'-chloro[1,1'-biphenyl]-3-yl)propoxy]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)  
FS 3D CONCORD  
MF C27 H28 Cl N O6 S  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



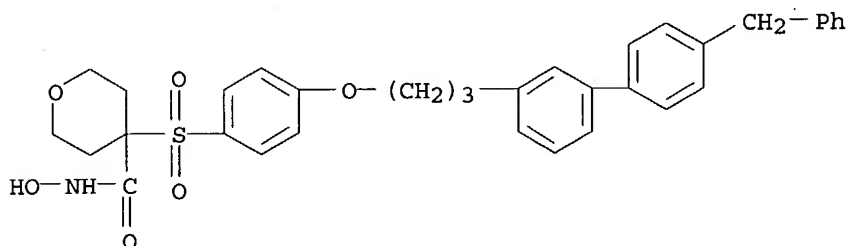
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 110 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 476186-79-5 REGISTRY  
CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[3-[4'-(phenylmethyl)[1,1'-biphenyl]-3-yl]propoxy]phenyl]sulfonyl]- (9CI) (CA INDEX NAME)  
FS 3D CONCORD  
MF C34 H35 N O6 S  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



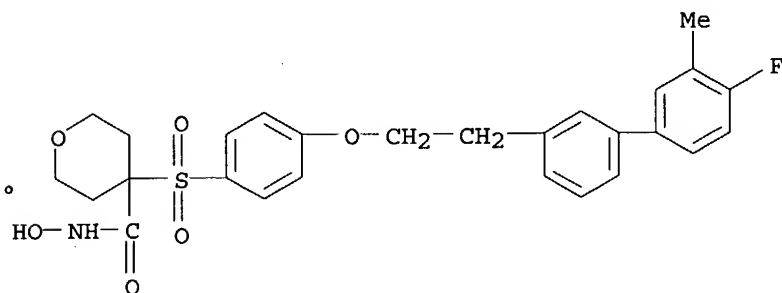
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2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 120 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 476186-69-3 REGISTRY  
CN 2H-Pyran-4-carboxamide, 4-[[4-[2-(4'-fluoro-3'-methyl[1,1'-biphenyl]-3-yl)ethoxy]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)  
FS 3D CONCORD  
MF C27 H28 F N O6 S  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



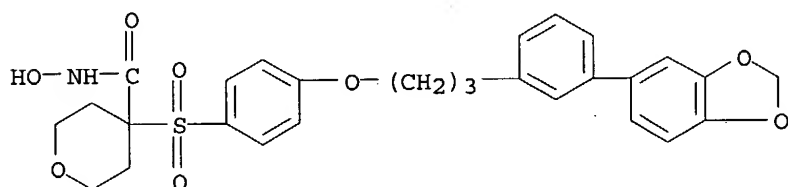
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2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 130 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 476186-59-1 REGISTRY  
CN 2H-Pyran-4-carboxamide, 4-[[4-[3-[3-(1,3-benzodioxol-5-yl)phenyl]propoxy]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)  
FS 3D CONCORD  
MF C28 H29 N O8 S  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



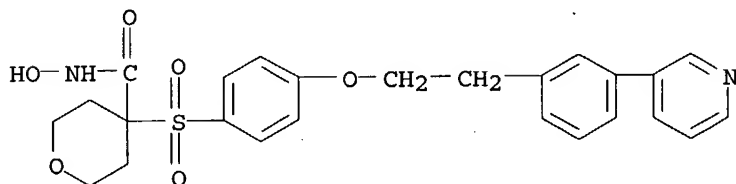
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2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 140 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 476186-49-9 REGISTRY  
CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[2-[3-(3-pyridinyl)phenyl]ethoxy]phenyl]sulfonyl]- (9CI) (CA INDEX NAME)  
FS 3D CONCORD  
MF C25 H26 N2 O6 S  
CI COM  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



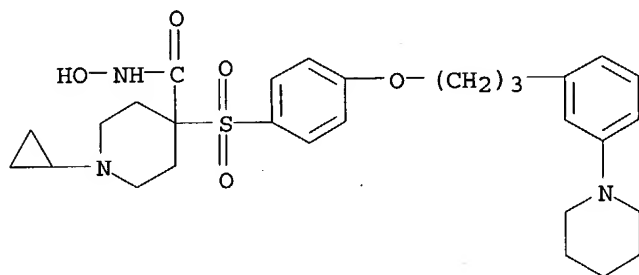
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2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 150 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
 RN 476186-39-7 REGISTRY  
 CN 4-Piperidinecarboxamide, 1-cyclopropyl-N-hydroxy-4-[[4-[3-[3-(1-piperidinyl)phenyl]propoxy]phenyl]sulfonyl]- (9CI) (CA INDEX NAME)  
 FS 3D CONCORD  
 MF C29 H39 N3 O5 S  
 CI COM  
 SR CA  
 LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



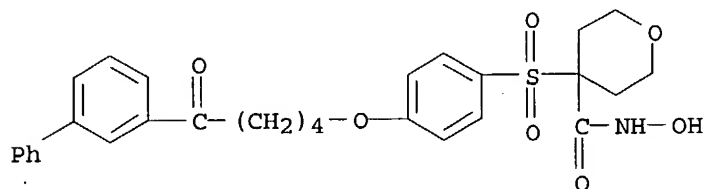
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2 REFERENCES IN FILE CA (1907 TO DATE)  
 2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747

L7 ANSWER 157 OF 157 REGISTRY COPYRIGHT 2004 ACS on STN  
 RN 476182-38-4 REGISTRY  
 CN 2H-Pyran-4-carboxamide, 4-[[4-[(5-[1,1'-biphenyl]-3-yl-5-oxopentyl)oxy]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)  
 FS 3D CONCORD  
 MF C29 H31 N O7 S  
 SR CA  
 LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

2 REFERENCES IN FILE CA (1907 TO DATE)  
 2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:163704

REFERENCE 2: 137:384747